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<input type="checkbox"/>	L1	(mur or murc or mur-c).clm.	20
<input type="checkbox"/>	L2	azzolina.in. or el-sherbeini.in. or elsherbeini.in.	29
<input type="checkbox"/>	L3	L2 and pseudomonas	6

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L2 and pseudomonas			6	

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Search Results - Record(s) 1 through 6 of 6 returned.

☐ 1. Document ID: WO 9961050 A1

Using default format because multiple data bases are involved.

L3: Entry 1 of 6

File: EPAB

Dec 2, 1999

PUB-NO: WO009961050A1

DOCUMENT-IDENTIFIER: WO 9961050 A1

TITLE: MURD PROTEIN AND GENE OF PSEUDOMONAS AERUGINOSA

PUBN-DATE: December 2, 1999

INVENTOR-INFORMATION:

NAME

COUNTRY

EL-SHERBEINI, MOHAMED

US

AZZOLINA, BARBARA

US

INT-CL (IPC): A61 K 39/00; A61 K 39/02; A61 K 39/108; C07 H 21/02; C07 H 21/04; C12 N 15/00; C12 N 15/20; C12 P 21/06; C12 P 21/04

EUR-CL (EPC): C12N009/00

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Abstract	Claims	EMC	Draw D
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☐ 2. Document ID: WO 200125251 A1, EP 1222197 A1

L3: Entry 2 of 6

File: DWPI

Apr 12, 2001

DERWENT-ACC-NO: 2001-308221

DERWENT-WEEK: 200254

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TITLE: New MraY gene and enzyme of Pseudomonas aeruginosa, useful in vitro assays for screening antibacterial compounds that target cell wall biosynthesis, particularly for screening antibiotics against Pseudomonads

INVENTOR: AZZOLINA, B; EL-SHERBEINI, M

PRIORITY-DATA: 1999US-157580P (October 4, 1999)

PATENT-FAMILY:

PUB-NO	PUB-DATE	LANGUAGE	PAGES	MAIN-IPC
<u>WO 200125251 A1</u>	April 12, 2001	E	022	C07H021/04
<u>EP 1222197 A1</u>	July 17, 2002	E	000	C07H021/04

INT-CL (IPC): C07 H 21/04; C12 N 15/00; C12 N 15/09; C12 P 21/06

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw. De
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☐ 3. Document ID: WO 200119979 A1

L3: Entry 3 of 6

File: DWPI

Mar 22, 2001

DERWENT-ACC-NO: 2001-281522

DERWENT-WEEK: 200129

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TITLE: New Pseudomonas aeruginosa MurC enzyme involved in bacterial cell wall biosynthesis, useful for identifying inhibitors of enzyme which are active against both gram positive and gram negative bacteria

INVENTOR: AZZOLINA, B; EL-SHERBEINI, M

PRIORITY-DATA: 1999US-154073P (September 14, 1999)

PATENT-FAMILY:

PUB-NO	PUB-DATE	LANGUAGE	PAGES	MAIN-IPC
<u>WO 200119979 A1</u>	March 22, 2001	E	039	C12N015/09

INT-CL (IPC): A61 K 31/70; C12 N 15/09

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw. De
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☐ 4. Document ID: WO 200119843 A1

L3: Entry 4 of 6

File: DWPI

Mar 22, 2001

DERWENT-ACC-NO: 2001-257872

DERWENT-WEEK: 200126

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TITLE: Novel purified and isolated Pseudomonas aeruginosa MurE polypeptide useful in assays to identify compounds that modulate activity of the polypeptide and for generation of antibodies against the polypeptide

INVENTOR: AZZOLINA, B; EL-SHERBEINI, M

PRIORITY-DATA: 1999US-154117P (September 15, 1999)

PATENT-FAMILY:

PUB-NO	PUB-DATE	LANGUAGE	PAGES	MAIN-IPC
<u>WO 200119843 A1</u>	March 22, 2001	E	041	C07H021/04

INT-CL (IPC): C07 H 21/04; C12 N 15/00; C12 N 15/09; C12 P 21/06

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWIC	Draw. De
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☐ 5. Document ID: WO 200118018 A1

L3: Entry 5 of 6

File: DWPI

Mar 15, 2001

DERWENT-ACC-NO: 2001-244554

DERWENT-WEEK: 200125

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TITLE: New Pseudomonas aeruginosa MurF polypeptide useful in assays to identify compounds that modulate activity of the polypeptide and for generation of antibodies against the polypeptide

INVENTOR: AZZOLINA, B; EL-SHERBEINI, M

PRIORITY-DATA: 1999US-153293P (September 10, 1999)

PATENT-FAMILY:

PUB-NO	PUB-DATE	LANGUAGE	PAGES	MAIN-IPC
WO 200118018 A1	March 15, 2001	E	040	C07H021/02

INT-CL (IPC): C07 H 21/02; C07 H 21/04; C12 N 1/14; C12 N 1/16; C12 N 1/18; C12 N 1/20; C12 N 5/00; C12 N 5/04; C12 N 5/10; C12 N 9/00; C12 N 15/00; C12 N 15/09; C12 N 15/63; C12 N 15/70; C12 N 15/74; G01 N 33/53

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Summary	Claims	KWC	Draw. D.
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☐ 6. Document ID: WO 9961050 A1, JP 2002516076 W, EP 1079855 A1

L3: Entry 6 of 6

File: DWPI

Dec 2, 1999

DERWENT-ACC-NO: 2000-072548

DERWENT-WEEK: 200239

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TITLE: New nucleic acid encoding the MurD protein of Pseudomonas aeruginosa, used to identify specific inhibitors

INVENTOR: AZZOLINA, B; EL-SHERBEINI, M

PRIORITY-DATA: 1998US-087308P (May 29, 1998)

PATENT-FAMILY:

PUB-NO	PUB-DATE	LANGUAGE	PAGES	MAIN-IPC
WO 9961050 A1	December 2, 1999	E	035	A61K039/00
JP 2002516076 W	June 4, 2002		040	C12N015/09
EP 1079855 A1	March 7, 2001	E	000	A61K039/00

INT-CL (IPC): A61 K 38/00; A61 K 39/00; A61 K 39/02; A61 K 39/108; A61 K 45/00; A61 P 31/04; C07 H 21/02; C07 H 21/04; C12 N 15/00; C12 N 15/09; C12 N 15/20; C12 P 21/04; C12 P 21/06

Full	Title	Citation	Front	Review	Classification	Date	Reference	Abstract	Summary	Claims	KWC	Draw. D.
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L1: Entry 2 of 20

File: PGPB

Mar 18, 2004

DOCUMENT-IDENTIFIER: US 20040052799 A1

TITLE: Nucleic acid and amino acid sequences relating to Helicobacter pylori for diagnostics and therapeutics

CLAIMS:

182. An isolated nucleic acid comprising a nucleotide sequence encoding an H. pylori pylori murC polypeptide or a fragment thereof, said nucleic acid comprising the nucleotide sequence of SEQ ID NO: 2845.

188. A purified H. pylori murC polypeptide or a fragment thereof, wherein said polypeptide comprises the amino acid sequence of SEQ ID NO: 7607.

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L1: Entry 2 of 20

File: PGPB

Mar 18, 2004

PGPUB-DOCUMENT-NUMBER: 20040052799

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20040052799 A1

TITLE: Nucleic acid and amino acid sequences relating to Helicobacter pylori for
diagnostics and therapeutics

PUBLICATION-DATE: March 18, 2004

INT-CL: [07] [A61 K 39/00](#), [A61 K 39/38](#)

US-CL-PUBLISHED: 424/184.1

US-CL-CURRENT: [424/184.1](#)[Previous Doc](#)[Next Doc](#)[Go to Doc#](#)

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L1: Entry 3 of 20

File: PGPB

Feb 12, 2004

DOCUMENT-IDENTIFIER: US 20040028702 A1

TITLE: Muramic acid derivative compounds

CLAIMS:

8. A library of compounds prepared by the method of claim 7 for screening for inhibiting Mur enzymes.

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L1: Entry 3 of 20

File: PGPB

Feb 12, 2004

PGPUB-DOCUMENT-NUMBER: 20040028702

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20040028702 A1

TITLE: Muramic acid derivative compounds

PUBLICATION-DATE: February 12, 2004

INT-CL: [07] [A61 K 39/02](#), [C08 B 37/00](#), [A61 K 31/739](#)US-CL-PUBLISHED: [424/234.1](#); [514/54](#), [536/53](#)US-CL-CURRENT: [424/234.1](#); [514/54](#), [536/53](#)[Previous Doc](#)[Next Doc](#)[Go to Doc#](#)

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L1: Entry 10 of 20

File: PGPB

Feb 7, 2002

PGPUB-DOCUMENT-NUMBER: 20020015678
PGPUB-FILING-TYPE: new
DOCUMENT-IDENTIFIER: US 20020015678 A1

TITLE: DIRECT ADSORPTION SCINTILLATION ASSAY FOR MEASURING ENZYME ACTIVITY AND
ASSAYING BIOCHEMICAL PROCESSES

PUBLICATION-DATE: February 7, 2002

INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
YUAN, ZHENGYU	FREMONT	CA	US	
CHEN, ZHONG-XIAO	BELMONT	CA	US	

US-CL-CURRENT: 424/1.11; 422/102, 435/4, 435/7.1, 435/7.2, 436/501, 436/518,
436/537

CLAIMS:

What is claimed is:

- 1) A method for analyzing a sample comprising: a) providing a sample containing one or more molecular species, wherein at least one of the molecular species is capable of stimulating scintillation; b) providing a scintillating material, wherein the surface of the scintillating material adsorbs at least one of the molecular species via a general molecular property-based binding interaction between the molecular species and the scintillating material, and where the scintillating material can be stimulated to scintillate by at least one of the adsorbed molecular species, but is generally not stimulated to scintillate by any molecular species which is not adsorbed; c) measuring the scintillation emitted by the scintillating material.
- 2) The method of claim 1, wherein the number of molecular species provided is at least two, and where at least one of said molecular species has a presence of, an absence of, or a degree of general molecular property-based binding interaction with the scintillating material distinct from the remainder of the molecular species.
- 3) The method of claim 1, wherein the general molecular property-based binding interaction is selected from the group consisting of charge-charge interactions, dipole-charge interactions, dipole-dipole interactions and hydrophobic interactions.
- 4) The method of claim 2, wherein the presence of, the absence of, or the degree of general molecular property-based binding interaction with the scintillating material is due to a chemical or biochemical transformation of one of said molecular species into another of said molecular species, further comprising the step of determining the progress of or degree of completion of the molecular transformation.

- 5) The method of claim 1, wherein the scintillating material is selected from the group consisting of scintillating plastics and scintillating glasses.
- 6) The method of claim 1, wherein the scintillating material is a plastic doped with with a scintillant.
- 7) The method of claim 5, wherein the scintillating plastic is selected from the group consisting of polystyrene doped with at least one scintillating fluor and polyvinyltoluene doped with at least one scintillating fluor.
- 8) The method of claim 2, wherein at least one of the at least two molecular species species provided is a substrate for an enzyme-catalyzed reaction or a series of enzyme-catalyzed reactions, another of the at least two molecular species is a product of the enzyme-catalyzed reaction or series of enzyme-catalyzed reactions and and has a presence of, absence of, or degree of general molecular property-based binding affinity for the scintillating material distinct from that of the substrate, substrate, and where the difference in general molecular property-based binding affinity is a result of the enzyme-catalyzed reaction or series of enzyme-catalyzed reactions.
- 9) The method of claim 8, wherein the general molecular property-based binding affinity is due to the presence of positive charge, the absence of positive charge, the presence of negative charge, the absence of negative charge, the presence of a dipole moment, the absence of a dipole moment, the presence of hydrophobicity, or the absence of hydrophobicity.
- 10) The method of claim 8, wherein the enzyme catalyzed reaction is selected from the group consisting of kinase catalyzed reactions, lipase catalyzed reactions, phosphatase catalyzed reactions, protease catalyzed reactions, and tRNA transferase catalyzed reactions.
- 11) The method of claim 8, wherein the enzyme catalyzed reaction is selected from the group consisting of the reaction cascade or any portion thereof for the sequential synthesis of uridinediphosphate-N-acetylmurami- c acid pentapeptide catalyzed by the enzymes MurA, MurB, MurC, MurD, MurE, and MurF.
- 12) The method of claim 8, wherein the enzyme catalyzed reaction is that catalyzed by MurA.
- 13) The method of claim 8, wherein the enzyme catalyzed reaction is that catalyzed by MurB.
- 14) The method of claim 8, wherein the enzyme catalyzed reaction is that catalyzed by MurC.
- 15) The method of claim 8, wherein the enzyme catalyzed reaction is that catalyzed by MurD.
- 16) The method of claim 8, wherein the enzyme catalyzed reaction is that catalyzed by MurE.
- 17) The method of claim 8, wherein the enzyme catalyzed reaction is that catalyzed by MurF.
- 18) The method of claim 8, wherein the enzyme catalyzed reaction is the reaction cascade for the sequential synthesis of uridinediphosphate-N-ace- tylmuramic acid pentapeptide catalyzed by the enzymes MurA, MurB, MurC, MurD, MurE, and MurF.

19) The method of claim 4, further comprising performing the method on a plurality of samples to effect a high throughput screen.

20) The method of claim 19, wherein the high throughput screen is used to identify compounds which inhibit an enzyme catalyzed reaction selected from the group consisting of the reaction cascade or any portion thereof for the sequential synthesis of uridinediphosphate-N-acetylmuramic acid pentapeptide catalyzed by the enzymes MurA, MurB, MurC, MurD, MurE, and MurF; kinase catalyzed reactions, lipase catalyzed reactions, phosphatase catalyzed reactions, protease catalyzed reactions, and tRNA transferase catalyzed reactions

21) A plate suitable for a direct adsorption binding assay, said plate comprised of a scintillating material and having one or more wells.

22) A plate suitable for a direct adsorption binding assay, said plate comprising wells coated with a scintillant material.

23) The plate of claim 21, wherein said wells are derivatized such that the walls of the wells are positively charged.

24) The plate of claim 21, wherein said wells are derivatized such that the walls of the wells are negatively charged.

25) The plate of claim 21, wherein said wells are derivatized such that the walls of the wells are hydrophobic.

26) The plate of claim 23, wherein the walls of the wells are derivatized with methyltrioctylammonium bromide.

27) The plate of claim 24, wherein the walls of the wells are derivatized with octadecyl sulfate.

28) The plate of claim 25, where the walls of the wells are derivatized with polylysine-N.sup.68-palmitate.

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L1: Entry 14 of 20

File: USPT

Apr 6, 1999

US-PAT-NO: 5891621

DOCUMENT-IDENTIFIER: US 5891621 A

TITLE: Metabolic pathway assay

DATE-ISSUED: April 6, 1999

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Chabin; Renee M.	Neptune	NJ		
Kuo; David W.	Princeton	NJ		
O'Connell; John F.	Cranbury	NJ		
Pompliano; David L.	Lawrenceville	NJ		
Wong; Kenny K.	Edison	NJ		

ASSIGNEE-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY	TYPE CODE
Merck & Co., Inc.	Rahway	NJ			02

APPL-NO: 08/ 936646 [\[PALM\]](#)

DATE FILED: September 24, 1997

INT-CL: [06] [C12 Q 1/00](#), [C12 Q 1/37](#), [C12 Q 1/18](#), [C12 Q 1/48](#)

US-CL-ISSUED: 435/4; 435/23, 435/24, 435/32, 435/15, 435/21, 435/7.91, 435/18, 435/16

US-CL-CURRENT: [435/4](#); [435/15](#), [435/16](#), [435/18](#), [435/21](#), [435/23](#), [435/24](#), [435/32](#), [435/7.91](#)

FIELD-OF-SEARCH: 435/4, 435/16, 435/23, 435/24, 435/32, 435/15, 435/21, 435/7.91, 435/18

PRIOR-ART-DISCLOSED:

OTHER PUBLICATIONS

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Tanner, et al., "Phosphinate Inhibitors of the d-Glutamic Acid-Adding Enzyme of Peptidoglycan Biosynthesis," J. Org. Chem., vol. 61, pp. 1756-1760 (1996).

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Jin, et al., "Structural Studies of Escherichia coli UDP-N-Acetylmuramate:L-Alanine Ligase," Biochemistry, vol. 35, pp. 1423-1431 (1996).

Auger, et al., "Synthesis of N-Acetylmuranic Acid Derivatives as Potential

Inhibitors of the D-Glutamic Acid-Adding Enzyme," J. prakt. Chem., vol. 337, pp. 351-357 (1995).

Pratviel-Sosa, et al., "Over-production, purification and properties of the uridine diphosphate N-acetylmuramoyl-L-Alanine: D-glutamate ligase from Escherichia coli," Eur. J. Bio. Chem., vol. 202, pp. 1169-1176 (1991).

Michaud, et al., "Partial purification and specificity studies of the D-glutamate-adding and d-alanyl-D-alanine-adding enzymes from Escherichia coli K12," Eur. J. Biochem., vol. 166, pp. 631-637 (1987).

LeRoux et al., "Synthesis of new peptide inhibitors of the meso-diaminopimelate-adding enzyme," Eur. J. Med. Chem., vol. 27, pp. 899-907 (1992).

Pratviel-Sosa, et al., "Effect of various analogues of D-glutamic acid on the D-glutamate-adding enzyme from Escherichia coli," FEMS, Microbiol. Letters, vol. 115, pp. 223-228 (1994).

Michaud et al., "Over-production, purification and properties of the uridine-diphosphate-N-acetylmuramoyl-L-alanyl-D-glutamate: meso-2,6-diaminopimelate ligase from Escherichia coli," Eur. J. Biochem. vol. 194, pp. 853-861 (1990).

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Abo-Ghaila, et al., "Specificity of the uridine-diphosphate-N-acetylmuramyl-L-alanyl-D-glutamate:meso-2, 6-diaminopimelate synthetase from Escherichia coli," Eur. J. Biochem., vol. 53, pp. 81-87 (1985).

Bugg and Walsh, "Intracellular Steps of Bacterial Cell Wall Peptidoglycan Biosynthesis: Enzymology, Antibiotics, and Antibiotic Resistance," Natural Product Reports (1992).

Presentation by Dr. D. Pompliano, Apr. 23, 1996, to University of Minnesota Departmental Staff Meeting.

Hakes & Dixon "New Vectors for High Level Expression of Recombinant Proteins in Bacteria," Anal. Biochem., vol. 202, pp. 293-298 (1992).

Reddy, et al., Mechanistic Analysis of UDP-N-acetylmuramyl:L-alanine Ligase from Escherichia coli presented at Enzyme Mechanism 15th Conference, Naples, Florida, Jan. 4, 1997-Jan. 8, 1997.

ART-UNIT: 163

PRIMARY-EXAMINER: Leary; Louise N.

ATTY-AGENT-FIRM: Fitch; Catherine D. Winokur; Melvin

ABSTRACT:

An in vitro screening assay which identifies enzyme inhibitors and allows for the simultaneous assay of many enzymes. Enzyme, substrate, co-factor, etc. concentrations are optimized so that inhibitors of any one of the enzymes in the pathway are equally likely to be detected. Necessarily, the flux of substrate through each enzyme should be nearly the same during the assay, i.e., each of the enzyme catalyzed steps must be equally rate-limiting. Preferably, optimal assay conditions are predicted by computer modeling. Further, the pathway conditions are optimized through variation of enzyme, starting substrate, co-substrate and co-factor concentrations. A positive response is initially detected as a change in the amount of the product generated at the end of the enzyme cascade as compared to a standard. A sample producing a positive result can be deconvoluted.

37 Claims, 2 Drawing figures

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L1: Entry 14 of 20

File: USPT

Apr 6, 1999

US-PAT-NO: 5891621

DOCUMENT-IDENTIFIER: US 5891621 A

TITLE: Metabolic pathway assay

DATE-ISSUED: April 6, 1999

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Chabin; Renee M.	Neptune	NJ		
Kuo; David W.	Princeton	NJ		
O'Connell; John F.	Cranbury	NJ		
Pompliano; David L.	Lawrenceville	NJ		
Wong; Kenny K.	Edison	NJ		

US-CL-CURRENT: [435/4](#); [435/15](#), [435/16](#), [435/18](#), [435/21](#), [435/23](#), [435/24](#), [435/32](#),
[435/7.91](#)

CLAIMS:

What is claimed is:

1. A kit which is comprised of an enzyme cascade comprising a first enzyme, a second enzyme and a substrate for the first enzyme.
2. The kit according to claim 1 wherein the first enzyme is suitable for changing the substrate for the first enzyme into a substrate for the second enzyme, and the second enzyme is suitable for changing the substrate for the second enzyme into a product of the second enzyme.
3. The kit according to claim 1 wherein the concentration of each component in the cascade is optimized to maintain uniform flux of substrate through the cascade.
4. The kit according to claim 1 additionally comprising one or more co-substrates for the first and second enzymes.
5. The kit according to claim 4 wherein the first enzyme is suitable for changing the substrate for the first enzyme into a substrate for the second enzyme, and the second enzyme is suitable for changing the substrate for the second enzyme into a product of the second enzyme.
6. The kit according to claim 5 wherein the concentration of each component in the cascade is optimized to maintain uniform flux of substrate through the cascade.

7. The kit according to claim 4 additionally comprising a third enzyme, a fourth enzyme, and one or more co-substrates for the third and fourth enzymes.
8. The kit according to claim 7 wherein the first enzyme is suitable for changing the substrate for the first enzyme into a substrate for the second enzyme; the second enzyme is suitable for changing the substrate for the second enzyme into a substrate for the third enzyme; the third enzyme is suitable for changing the substrate for the third enzyme into a substrate for the fourth enzyme; and the fourth enzyme is suitable for changing the substrate for the fourth enzyme into a product of the fourth enzyme.
9. The kit according to claim 8 wherein the concentration of each enzyme in the cascade is optimized to maintain uniform flux of substrate through the cascade.
10. The kit according to claim 9 wherein the first enzyme is the gene product of murC, the second enzyme is the gene product of murD, the third enzyme is the gene product of murE, and the fourth enzyme is the gene product of murF.
11. The kit according to claim 10 wherein
- the substrate for the first enzyme is uridyl-5'-diphosphate N-acetyl muramic acid;
- the co-substrates for the first enzyme are L-alanine and adenosine 5'-triphosphate;
- the substrate for the second enzyme is uridyl-5'-diphosphate N-acetyl muramyl-L-alanine;
- the co-substrates for the second enzyme are D-glutamic acid, and adenosine 5'-triphosphate;
- the substrate for the third enzyme is uridyl-5'-diphosphate N-acetyl muramyl-L-alanyl-.gamma.-D-glutamic acid;
- the co-substrates for the third enzyme are meso-diaminopimelic acid and adenosine 5'-triphosphate;
- the substrate for the fourth enzyme is uridyl-5'-diphosphate N-acetyl muramyl-L-alanyl-.gamma.-D-glutamyl-meso-dipimelic acid; and
- the co-substrates for the fourth enzyme are D-alanyl-D-alanine and adenosine 5'-triphosphate.
12. The kit according to claim 7 additionally comprising a fifth enzyme, a sixth enzyme, and one or more co-substrates for the fifth and sixth enzymes.
13. The kit assay according to claim 12 wherein the first enzyme is suitable for changing the substrate for the first enzyme into a substrate for the second enzyme; the second enzyme is suitable for changing the substrate for the second enzyme into a substrate for the third enzyme; the third enzyme is suitable for changing the substrate for the third enzyme into a substrate for the fourth enzyme; the fourth enzyme is suitable for changing the substrate for the fourth enzyme into a substrate for the fifth enzyme; the fifth enzyme

is suitable for changing the substrate for the fifth enzyme into a substrate for the sixth enzyme; and the sixth enzyme is suitable for changing the substrate for the sixth enzyme into a product of the sixth enzyme.

14. The kit according to claim 13 wherein the concentration of each enzyme in the cascade is optimized to maintain uniform flux of substrate through the cascade.

15. The kit according to claim 14 wherein the first enzyme is the gene product of *murA*, the second enzyme is the gene product of *murB*, the third enzyme is the gene product of *murC*, the fourth enzyme is the gene product of *murD*, the fifth enzyme is the gene product of *MurE*, and the sixth enzyme is the product of *MurF*.

16. The kit according to claim 15 wherein

the substrate for the first enzyme is uridyl-5'-diphosphate-N-acetyl-glucosamine;

the co-substrate for the first enzyme is phosphoenolpyruvate;

the substrate for the second enzyme is uridyl-5'-diphosphate-N-acetyl-enolpyruvylglucosamine;

the co-substrates for the second enzyme are nicotinamide adenine dinucleotide phosphate reduced form and flavin adenine dinucleotide;

the substrate for the third enzyme is uridyl-5'-diphosphate N-acetyl muramic acid;

the co-substrates for the third enzyme are L-alanine and adenosine 5'-triphosphate;

the substrate for the fourth enzyme is uridyl-5'-diphosphate N-acetyl muramyl-L-alanine;

the co-substrates for the fourth enzyme are D-glutamic acid, and adenosine 5'-triphosphate;

the substrate for the fifth enzyme is uridyl-5'-diphosphate N-acetyl muramyl-L-alanyl-.gamma.-D-glutamic acid;

the co-substrates for the fifth enzyme are meso-diaminopimelic acid and adenosine 5'-triphosphate;

the substrate for the sixth enzyme is uridyl-5'-diphosphate N-acetyl muramyl-L-alanyl-.gamma.-D-glutamyl-meso-dipimelic acid; and

the co-substrates for the sixth enzyme are D-alanyl-D-alanine and adenosine 5'-5'-triphosphate.

17. A method of screening for a biologically active compound which comprises:

combining an enzyme cascade, comprising a first enzyme, a second enzyme and a substrate for the first enzyme, with a compound suspected of having biological

activity; and

measuring the concentration of the products of the enzymes and comparing to a standard.

18. The method according to claim 17 wherein the concentrations of products are measured by radio-labeled HPLC.

19. The method according to claim 17 wherein the first enzyme is suitable for changing the substrate for the first enzyme into a substrate for the second enzyme, and the second enzyme is suitable for changing the substrate for the second enzyme into a product of the second enzyme.

20. The method according to claim 19 wherein the concentration of each enzyme in the cascade is optimized to maintain uniform flux of substrate through the cascade.

21. The method according to claim 17 wherein the enzyme cascade additionally comprises one or more co-substrates for the first and second enzymes.

22. The method according to claim 21 wherein the concentrations of products are measured by radio-labeled HPLC.

23. The method according to claim 21 wherein the first enzyme is suitable for changing the substrate for the first enzyme into a substrate for the second enzyme, and the second enzyme is suitable for changing the substrate for the second enzyme into a product of the second enzyme.

24. The method according to claim 23 wherein the concentration of each enzyme in the cascade is optimized to maintain uniform flux of substrate through the cascade.

25. The method according to claim 21 wherein the enzyme cascade additionally comprises a third enzyme, a fourth enzyme and one or more co-substrates for the the third and fourth enzymes.

26. The method according to claim 25 wherein the concentrations of products are measured by radio-labeled HPLC.

27. The method according to claim 25 wherein the first enzyme is suitable for changing the substrate for the first enzyme into a substrate for the second enzyme; the second enzyme is suitable for changing the substrate for the second enzyme into a substrate for the third enzyme; the third enzyme is suitable for changing the substrate for the third enzyme into a substrate for the fourth enzyme; and the fourth enzyme is suitable for changing the substrate for the fourth enzyme into a product of the fourth enzyme.

28. The method according to claim 27 wherein the concentration of each enzyme in the cascade is optimized to maintain uniform flux of substrate through the cascade.

29. The method according to claim 28 wherein the first enzyme is the gene product of murC, the second enzyme is the gene product of murD, the third enzyme is the gene product of murE, and the fourth enzyme is the gene product of murF.

30. The method according to claim 29 wherein

the substrate for the first enzyme is uridyl-5'-diphosphate N-acetyl muramic acid;

the co-substrates for the first enzyme are L-alanine and adenosine 5'-triphosphate;

the substrate for the second enzyme is uridyl-5'-diphosphate N-acetyl muramyl-L-alanine;

the co-substrates for the second enzyme are D-glutamic acid, and adenosine 5'-triphosphate;

the substrate for the third enzyme is uridyl-5'-diphosphate N-acetyl muramyl-L-alanyl-.gamma.-D-glutamic acid;

the co-substrates for the third enzyme are meso-diaminopimelic acid and adenosine 5'-triphosphate;

the substrate for the fourth enzyme is uridyl-5'-diphosphate N-acetyl muramyl-L-alanyl-.gamma.-D-glutamyl-meso-dipimelic acid; and

the co-substrates for the fourth enzyme are D-alanyl-D-alanine and adenosine 5'-triphosphate.

31. The method according to claim 25 wherein the enzyme cascade additionally comprises a fifth enzyme, a sixth enzyme, and one or more co-substrates for the the fifth and sixth enzymes.

32. The method according to claim 31 wherein the concentrations of products are measured by radio-labeled HPLC.

33. The method according to claim 31 wherein the first enzyme is suitable for changing the substrate for the first enzyme into a substrate for the second enzyme; the second enzyme is suitable for changing the substrate for the second enzyme into a substrate for the third enzyme; the third enzyme is suitable for changing the substrate for the third enzyme into a substrate for the fourth enzyme; the fourth enzyme is suitable for changing the substrate for the fourth enzyme into a substrate for the fifth enzyme; the fifth enzyme is suitable for changing the substrate for the fifth enzyme into a substrate for the sixth enzyme; and the sixth enzyme is suitable for changing the substrate for the sixth enzyme into a product of the sixth enzyme.

34. The method according to claim 33 wherein the concentration of each enzyme in the cascade is optimized to maintain uniform flux of substrate through the cascade.

35. The method according to claim 34 wherein the first enzyme is the gene product of murA, the second enzyme is the gene product of murB, the third enzyme is the gene product of murC, the fourth enzyme is the gene product of murD, the fifth enzyme is the gene product of murE, and the sixth enzyme is the product of murF.

36. The method according to claim 35 wherein

the substrate for the first enzyme is uridyl-5'-diphosphate-N-acetyl-glucosamine;

the co-substrate for the first enzyme is phosphoenolpyruvate;

the substrate for the second enzyme is uridyl-5'-diphosphate-N-acetyl-enolpyruvylglucosamine;

the co-substrates for the second enzyme are nicotinamide adenine dinucleotide phosphate reduced form and flavin adenine dinucleotide;

the substrate for the third enzyme is uridyl-5'-diphosphate N-acetyl muramic acid;

the co-substrates for the third enzyme are L-alanine and adenosine 5'-triphosphate;

the substrate for the fourth enzyme is uridyl-5'-diphosphate N-acetyl muramyl-L-alanine;

the co-substrates for the fourth enzyme are D-glutamic acid, and adenosine 5'-triphosphate;

the substrate for the fifth enzyme is uridyl-5'-diphosphate N-acetyl muramyl-L-alanyl-.gamma.-D-glutamic acid;

the co-substrates for the fifth enzyme are meso-diaminopimelic acid and adenosine 5'-triphosphate;

the substrate for the sixth enzyme is uridyl-5'-diphosphate N-acetyl muramyl-L-alanyl-.gamma.-D-glutamyl-meso-dipimelic acid; and

the co-substrates for the sixth enzyme are D-alanyl-D-alanine and adenosine 5'-5'-triphosphate.

37. A high-throughput in vitro screening method for detecting a biologically active compound which is comprised of:

(a) combining an enzyme cascade comprising a first enzyme, a second enzyme, a third enzyme, a fourth enzyme, a fifth enzyme and a sixth enzyme, and a labeled substrate for the first enzyme with a compound suspected of having biological activity; wherein:

the first enzyme is the gene product of murA, the second enzyme is the gene product of murB, the third enzyme is the gene product of murC, the fourth enzyme is the gene product of murD, the fifth enzyme is the gene product of murE, and the sixth enzyme is the product of murF; and

(b) measuring the concentrations of the products of the enzymes and comparing to a standard, by absorbing the product of the sixth enzyme onto resin and detecting the amount of label and comparing the amount of label to control.

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- ☐ 1. [6838255](#). 23 Jun 98; 04 Jan 05. MurC. Burnham; Martin Karl Russell, et al. 435/69.1; 435/252.3 435/252.33 435/320.1 536/23.7. C12P021/02.
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(mur or murc or mur-c).ti,ab,clm. not L1 not L3	15

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<input type="checkbox"/>	L3	UDP-N-acetylmuramoyl	2
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MURC_PSEAE (Q9HW02)

UDP-N-acetylmuramate--L-alanine ligase (EC 6.3.2.8) (UDP-N-acetylmuramoyl-L-alanine synthetase). {GENE: Name=murC; OrderedLocusNames=PA4411} - Pseudomonas aeruginosa

MURC_PSEPK (Q88N75)

UDP-N-acetylmuramate--L-alanine ligase (EC 6.3.2.8) (UDP-N-acetylmuramoyl-L-alanine synthetase). {GENE: Name=murC; OrderedLocusNames=PP1338} - Pseudomonas putida (strain KT2440)

MURC_PSESM (Q87WY6)

UDP-N-acetylmuramate--L-alanine ligase (EC 6.3.2.8) (UDP-N-acetylmuramoyl-L-alanine synthetase). {GENE: Name=murC; OrderedLocusNames=PSPTO4407} - Pseudomonas syringae (pv. tomato)

MURC_RALSO (Q8XVI8)

UDP-N-acetylmuramate--L-alanine ligase (EC 6.3.2.8) (UDP-N-acetylmuramoyl-L-alanine synthetase). {GENE: Name=murC; OrderedLocusNames=RSc2843; ORFNames=RS00262} - Ralstonia solanacearum (Pseudomonas solanacearum)

MURC_RHOPA (P61682)

UDP-N-acetylmuramate--L-alanine ligase (EC 6.3.2.8) (UDP-N-acetylmuramoyl-L-alanine synthetase). {GENE: Name=murC; OrderedLocusNames=RPA3529} - Rhodopseudomonas palustris

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Q62GS8

UDP-N-acetylmuramate--alanine ligase (EC 6.3.2.8) {GENE:Name=murC;
OrderedLocusNames=BMA2550} - Burkholderia mallei (Pseudomonas mallei)

Q63QJ8

UDP-N-acetylmuramate--alanine ligase (EC 6.3.2.8) {GENE:Name=murC;
OrderedLocusNames=BPSL3024} - Burkholderia pseudomallei (Pseudomonas pseudomallei)



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tr|Q5PDC7                  -MNTQQ-----LAKLRSIVPEMRRVRHIHFVGIGGAGMGGLAEVLANEGYQISGSDLAPN
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PileUp

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sp Q88N75 MURC_PSEPK	PTVTYGFSEE	ADIRAINVRQ	QGMQTHFTVL	RRDREPLEVS	VNMPGNHNVL
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sp P65470 MURC_ECOL6	QTTTYGFSED	ADVRVEDYQQ	IGPQGHFTLL	RQDKEPMRVT	LNAPGRHNAL
tr Q5PDC7	QTTTYGFSED	ADVRVEDYQQ	IGPQGHFTLL	RQGMPLDHVT	LNAPGRHNAL
sp P57818 MURC_PASMU	QVITYGFSEK	ADYRIEDYQQ	TGFQGHYTVI	TPSGERIDVL	LNVPGRHNAL

sp Q88N75 MURC_PSEPK	NALATIAIAT	DEGITDEAIV	QGLSGFQGVG	RRFQVYGELP	VE..GGS...
sp Q87WY6 MURC_PSESM	NSLATIAIAT	DEGVSDAIV	QGLSGFQGVG	RRFQVYGELP	VE..GGH...

sp P65470 MURC_ECOL6	NAAAAVAVAT	EEGIDDEAIL	RALESFQGTG	RRFDLGEFP	LEPVNGKSGT
tr Q5PDC7	NAAAAVAVAT	EEGIDDDAIL	RALESFQGTG	RRFDLGEFP	LEPVNGKAGT
sp P57818 MURC_PASMU	NATAALVVAK	EEGIGNEAIL	AALADFQAG	RRFDQLGQF.	IRP.NGK...
sp Q88N75 MURC_PSEPK	VMLVDDYGGH	PTEVAAVIKA	VRGGWPSRRL	VIVYQPHRYS	RTRDLYDDFV
sp Q87WY6 MURC_PSESM	VMLVDDYGGH	PREVAAVISA	VRGGWPDRL	VMVYQPHRFS	RTRDLYDDFV
sp P65470 MURC_ECOL6	AMLVDDYGGH	PTEVDATIKA	ARAGWPDKNL	VMLFQPHRET	RTRDLYDDFA
tr Q5PDC7	AMLVDDYGGH	PTEVDATIKA	ARAGWPDKNL	VMLFQPHRYT	RTRDLYDDFA
sp P57818 MURC_PASMU	VMLVDDYGGH	PTEVGVTIQA	ARQGWENKRI	AMIFQPHRYS	RTRDLFDDFV
sp Q88N75 MURC_PSEPK	QVLGDANVLL	LMEVYPAGEE	PIPGADSRQL	CHSIRQRGKL	DPIYIERGAE
sp Q87WY6 MURC_PSESM	QVLAEANVLL	LMEVYPAGEE	PVPGADSRNL	CHSIRQRGQL	DPIYIERGVE
sp P65470 MURC_ECOL6	NVLTQVDTL	MLEVYPAGEA	PIPGADSRSL	CRTIRGRGKI	DPILVPDPAQ
tr Q5PDC7	NVLTQVDALL	MLDVYPAGEA	PIPGADSRSL	CRTIRNRGKI	DPILVSDPAQ
sp P57818 MURC_PASMU	QVLSQVDVLI	MLDVYAAGET	PIAGADSRSL	CRSIRNLGKV	DPIFVSDHRQ
sp Q88N75 MURC_PSEPK	LAPLVKPLL	AGDILLCQGA	GDVGGLAPQL	MKSPLFAGAK	QEKSK....
sp Q87WY6 MURC_PSESM	LAPLVKPLL	AGDILLCQGA	GDIAGLAPRL	LNSPLFVGAK	VASTEGKLK
sp P65470 MURC_ECOL6	VAEMLAPVLT	GNDLILVQGA	GNIGKIARSL	AEIKLKPQTP	EEEQHD...
tr Q5PDC7	VATMLAPVLT	GNDLILVQGA	GNVGKIARYL	SEIKLKPQIQ	EEEQHG...
sp P57818 MURC_PASMU	LGEILDQVLQ	DGDLILAQGA	GNVSKIARQL	AETWTKE...

CLUSTAL W (1.74) multiple sequence alignment

```

tr|Q63QJ8      -----MKHIVKHIHFVIGGAGMSGIAEVLVNLGYQVSGSDLARNVTERLEAL
tr|Q62GS8      -----MKHIVKHIHFVIGGAGMSGIAEVLVNLGYQVSGSDLARNVTERLEAL
sp|Q9HW02|MURC_PSEAE  MVKEPNGVTRTMRIRRIHFVIGGAGMCGIAEVLNLGYEVSGSDLKASAVTERLEKF
                        : ::*****.*****:*****.***** :

tr|Q63QJ8      ARVSIGHDAANIEGANAVVSTAVRSDNPEVLAARRLRVPIVPRAVMLAELMRLKQGIA
tr|Q62GS8      ARVSIGHDAANIEGANAVVSTAVRSDNPEVLAARRLRVPIVPRAVMLAELMRLKQGIA
sp|Q9HW02|MURC_PSEAE  AQIFIGHQAENADGADVLVSSAINRANPEVASALERRI PVVPRAEMLAELMRYRHGIA
                        *: : **:* * :*:*:*:*:*:  **** :* . *:*:***** ***** :*:**

tr|Q63QJ8      AGTHGKTTTTSLVASVLAAGGLDPTFVIGGRLTSAGANARLGTGDFIVAEADESDASFL
tr|Q62GS8      AGTHGKTTTTSLVASVLAAGGLDPTFVIGGRLTSAGANARLGMGDFIVAEADESDASFL
sp|Q9HW02|MURC_PSEAE  AGTHGKTTTTSLIASVFAAGGLDPTFVIGGRLNAAGTNAQLGASRYLVAEADESDASFL
                        *****;***:*****.***:***:*** . :*****

tr|Q63QJ8      LYPVIEVITNIDADHMDTYGHDFARLKQAFIEFTQRLPFYGSAAVCIDDANVRQIVPLI
tr|Q62GS8      LYPVIEVITNIDADHMDTYGHDFARLKQAFIEFTQRLPFYGSAAVCIDDANVRQIVPLI
sp|Q9HW02|MURC_PSEAE  LQPMVAVVTNIDADHMATYGGDFNKLKKT FVEFLHNL PFYGLAVMCVDDPVVREILPQI
                        * *: : *:***** ** * :*:*:*:* .***** *:*:*. *:*:**

tr|Q63QJ8      KPVVRYGFAADAQVRAENVEARDGRMHFTVRREGREPLPVVLNLPGLHNVQNALAAIAI
tr|Q62GS8      KPVVRYGFAADAQVRAENVEARDGRMHFTVRREGREPLPVVLNLPGLHNVQNALAAIAI
sp|Q9HW02|MURC_PSEAE  RPTVTYGLSEDAADVRAINIRQEGMRTWFTVLRPEREPLDVSVNMPGLHNVLSLATIVI
                        :*. * *: : **:*** *: . . * *** * **** * *:***** *:*:*.

tr|Q63QJ8      TDLDVADAAIQQALAEFNGVGRRFQRYGEIAAAGGGAYTLIDDYGHHPVEMAATIAAAR
tr|Q62GS8      TDLDVADAAIQQALAEFNGVGRRFQRYGEIAAAGGGAYTLIDDYGHHPVEMAATIAAAR
sp|Q9HW02|MURC_PSEAE  TDEGISDEAIVQGLSGFQGVGRRFQVYGE LQVEGG-SVMLVDDYGHHPREVA AVIKAIR
                        ** .*: * ** *: : *:***** ***: . ** : *:***** *:*. * *

tr|Q63QJ8      AFPGRRLVLAFQPHRYTRTRDCFDDFVNVLSTVDALVLTEVYAAGEAPISTANGDALSR
tr|Q62GS8      AFPGRRLVLAFQPHRYTRTRDCFDDFVNVLSTVDALVLTEVYAAGEAPISTANGDALSR
sp|Q9HW02|MURC_PSEAE  GWPERRLVMVYQPHRYTRTRDLYEDFVQVLGEANVLLLMEVYPAGEEPIPGADSRQLCH
                        .:* ****: :***** :*:*:*. .*: * **.* ** .*: .*:

tr|Q63QJ8      LRAAGKVEPVFVATVDEVPDALAKLARDGDVVITMGAGSIGGVPGKLAQDTQQKG----
tr|Q62GS8      LRAAGKVEPVFVATVDEVPDALAKLARDGDVVITMGAGSIGGVPGKLAQDTQQKG----
sp|Q9HW02|MURC_PSEAE  IRQRGQLDPIYFERDADLAPLVKPLL RAGDILL CQAGADVGG LAPQLIKNPLFAGKGGK
                        :* *: :*:*. .: . : * * *: : : ***:***. :* :. *

tr|Q63QJ8      -
tr|Q62GS8      -
sp|Q9HW02|MURC_PSEAE  A

```

FileUp

MSF: 481 Type: P Check: 5274 ..

Name: tr|Q63QJ8 oo Len: 481 Check: 8421 Weight: 0.100

Name: tr|Q62GS8 oo Len: 481 Check: 8078 Weight: 0.100

Name: sp|Q9HW02|MURC_PSEAE oo Len: 481 Check: 8775 Weight: 0.100

//

tr Q63QJ8	MKHIVKHIHF	VGIGGAGMSG	IAEVLVNLGY	QVSGSDLARN
tr Q62GS8	MKHIVKHIHF	VGIGGAGMSG	IAEVLVNLGY	QVSGSDLARN
sp Q9HW02 MURC_PSEAE		MVKEPNGVTR	TMRRIRRIHF	VGIGGAGMCG	IAEVLLNLGY

tr Q63QJ8		AVTERLEALG	ARVSIGHDAA	NIEGANAVVV	STAVRSDNPE	VLAARRLRVP
tr Q62GS8		AVTERLEALG	ARVSIGHDAA	NIEGANAVVV	STAVRSDNPE	VLAARRLRVP
sp Q9HW02 MURC_PSEAE		AVTERLEKFG	AQIFIGHQAE	NADGADVLVV	SSAINRANPE	VASALERRIP

tr Q63QJ8		IVPRAVMLAE	LMRLKQGIAI	AGTHGKTTT	SLVASVLAAG	GLDPTFVIGG
tr Q62GS8		IVPRAVMLAE	LMRLKQGIAI	AGTHGKTTT	SLVASVLAAG	GLDPTFVIGG
sp Q9HW02 MURC_PSEAE		VVPRAEMLAE	LMRYRHGIAV	AGTHGKTTT	SLIASVFAAG	GLDPTFVIGG

tr Q63QJ8		RLTSAGANAR	LGTGDFIVAE	ADESASFLN	LYPVIEVITN	IDADHMDTYG
tr Q62GS8		RLTSAGANAR	LGMGDFIVAE	ADESASFLN	LYPVIEVITN	IDADHMDTYG
sp Q9HW02 MURC_PSEAE		RLNAAGTNAQ	LGASRYLVAE	ADESASFLH	LQPMVAVVTN	IDADHMDTYG

tr Q63QJ8		HDFARLKQAF	IEFTQRLPFY	GSAVVCIDDA	NVRQIVPLIS	KPVVRYGFAA
tr Q62GS8		HDFARLKQAF	IEFTQRLPFY	GSAVVCIDDA	NVRQIVPLIS	KPVVRYGFAA
sp Q9HW02 MURC_PSEAE		GDFNKLKKT	VEFLHNLPHY	GLAVMCVDDP	VVREILPQIA	RPTVTYGLSE

tr Q63QJ8		DAQVRAENVE	ARDGRMHFTV	RREGREPLPV	VLNLPGLHNV	QNALAAIAIA
tr Q62GS8		DAQVRAENVE	ARDGRMHFTV	RREGREPLPV	VLNLPGLHNV	QNALAAIAIA
sp Q9HW02 MURC_PSEAE		DADVRAINIR	QEGMRTWFTV	LRPEREPLDV	SVNMPGLHNV	LNSLATIVIA

tr Q63QJ8		TDLDVADAAI	QQALAEFNGV	GRRFQRYGEI	AAAGGGAYTL	IDDYGHHPVE
tr Q62GS8		TDLDVADAAI	QQALAEFNGV	GRRFQRYGEI	AAAGGGAYTL	IDDYGHHPVE
sp Q9HW02 MURC_PSEAE		TDEGISDEAI	VQGLSGFQGV	GRRFQVYGEI	QVEGG.SVML	VDDYGHHPRE

tr Q63QJ8		MAATIAAARG	AFPGRRLVLA	FQPHRYTRTR	DCFDDFVNVL	STVDALVLTE
tr Q62GS8		MAATIAAARG	AFPGRRLVLA	FQPHRYTRTR	DCFDDFVNVL	STVDALVLTE
sp Q9HW02 MURC_PSEAE		VAAVIKAIRG	GWPERRLVMV	YQPHRYTRTR	DLYEDFVQVL	GEANVLLLME

tr Q63QJ8		VYAAGEAPIS	TANGDALSRA	LRAAGKVEPV	FVATVDEVPD	ALAKLARDGD
tr Q62GS8		VYAAGEAPIS	TANGDALSRA	LRAAGKVEPV	FVATVDEVPD	ALAKLARDGD
sp Q9HW02 MURC_PSEAE		VYPAGEEPIP	GADSRQLCHS	IRQRGQLDPI	YFERDADLAP	LVKPLLKRGD

tr Q63QJ8		VVITMGAGSI	GGVPGKLAQD	TQQKG.....	
-----------	--	------------	------------	------------	--

tr Q62GS8	VVITMGAGSI	GGVPGKLAQD	TQQKG.....
sp Q9HW02 MURC_PSEAE	ILLCQGAGDV	GGLAPQLIKN	PLFAGKGGKG A

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L1: Entry 1 of 1

File: USPT

Apr 3, 2001

US-PAT-NO: 6211161

DOCUMENT-IDENTIFIER: US 6211161 B1

TITLE: UDP-N-acetylmuramoyl-l-alanine:D-glutamate ligase (MURD) of staphylococcus aureus

DATE-ISSUED: April 3, 2001

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Beattie; David T	Boston	MA		
Deresiewicz; Robert L	Boston	MA		
Lowe; Adrian M	Brighton	MA		
Nicholas; Richard O	Collegeville	PA		
Palmer; Leslie Marie	Audubon	PA		
Pratt; Julie M	Wigston			GB
Lonetto; Michael A	Collegeville	PA		
Hodgson; John E	Paris			FR

US-CL-CURRENT: 514/44; 435/252.3, 435/254.11, 435/320.1, 435/325, 435/455, 435/471, 435/69.1, 536/23.1, 536/23.2, 536/23.4, 536/23.7

CLAIMS:

What is claimed is:

1. An isolated polynucleotide comprising a first polynucleotide or the full complement of the entire length of the first polynucleotide, wherein the first polynucleotide comprises SEQ ID NO:1.
2. A vector comprising the isolated polynucleotide of claim 1.
3. An isolated host cell comprising the vector of claim 2.
4. A process for producing a polypeptide comprising the step of culturing the host cell of claim 3 under conditions sufficient for the production of the polypeptide, wherein the polypeptide is encoded by the first polynucleotide.
5. The isolated polynucleotide of claim 1 encoding a fusion polypeptide, wherein the fusion polypeptide comprises SEQ ID NO:2.
6. An isolated polynucleotide comprising a first polynucleotide or the full complement of the entire length of the first polynucleotide, wherein the first polynucleotide encodes the same mature polypeptide, expressed by the MurD gene

contained in Staphylococcus aureus WCUH 29 contained in NCIMB Deposit No. 40771.

7. The isolated polynucleotide of claim 6 encoding a fusion polypeptide, wherein the first polynucleotide encodes part of the fusion polypeptide.

8. An isolated polynucleotide comprising a first polynucleotide or the full complement of the entire length of the first polynucleotide, wherein the first polynucleotide encodes a polypeptide comprising the amino acid sequence set forth in SEQ ID NO:2.

9. A vector comprising the isolated polynucleotide of claim 8.

10. An isolated host cell comprising the vector of claim 9.

11. A process for producing a polypeptide comprising the step of culturing the host cell of claim 10 under conditions sufficient for the production of the polypeptide, wherein the polypeptide is encoded by the first polynucleotide.

12. The isolated polynucleotide of claim 8 encoding a fusion polypeptide, wherein the fusion polypeptide comprises SEQ ID NO:2.

13. An isolated polynucleotide comprising a first polynucleotide or the full complement of the entire length of the first polynucleotide, wherein the first polynucleotide encodes a polypeptide consisting of the amino acid sequence set forth in SEQ ID NO:2.

14. A vector comprising the isolated polynucleotide of claim 13.

15. An isolated host cell comprising the vector of claim 14.

16. A process for producing a polypeptide comprising the step of culturing the host cell of claim 15 under conditions sufficient for the production of the polypeptide, wherein the polypeptide is encoded by the first polynucleotide.

17. A method for producing antibodies in a mammal comprising: delivering to a tissue of the mammal a nucleic acid vector to direct expression in vivo of a polypeptide from an isolated polynucleotide of claim 8, wherein the polypeptide polypeptide is effective to induce an immunological response to the amino acid sequence of SEQ ID NO:2; and, wherein the polypeptide is expressed in vivo and induces an immunological response to produce antibodies to the amino acid sequence of SEQ ID NO:2.

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L1: Entry 1 of 1

File: USPT

Apr 3, 2001

US-PAT-NO: 6211161

DOCUMENT-IDENTIFIER: US 6211161 B1

TITLE: UDP-N-acetylmuramoyl-l-alanine:D-glutamate ligase (MURD) of staphylococcus aureus

DATE-ISSUED: April 3, 2001

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Beattie; David T	Boston	MA		
Deresiewicz; Robert L	Boston	MA		
Lowe; Adrian M	Brighton	MA		
Nicholas; Richard O	Collegeville	PA		
Palmer; Leslie Marie	Audubon	PA		
Pratt; Julie M	Wigston			GB
Lonetto; Michael A	Collegeville	PA		
Hodgson; John E	Paris			FR

ASSIGNEE-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY	TYPE	CODE
Brigham & Women's Hospital	Boston	MA			02	
Virus Research Institute	Cambridge	MA			02	
SmithKline Beecham Corporation	Philadelphia	PA			02	
SmithKline Beecham plc,				GB	03	

APPL-NO: 09/ 147928 [\[PALM\]](#)

DATE FILED: March 19, 1999

PARENT-CASE:

RELATED APPLICATIONS This application claims benefit of U.S. Provisional Patent Application No. 60/061,064 filed Oct. 3, 1997.

PCT-DATA:

APPL-NO	DATE-FILED	PUB-NO	PUB-DATE	371-DATE	102(E)-DATE
PCT/US98/20773	October 2, 1998	WO99/17794	Apr 15, 1999	Mar 19, 1999	Mar 19, 1999

INT-CL: [07] [A61 K 48/00](#), [C07 H 21/00](#), [C12 N 15/31](#), [C12 N 15/52](#), [C12 N 15/63](#)

US-CL-ISSUED: 514/44; 435/69.1, 435/252.3, 435/254.11, 435/325, 435/320.1, 435/455, 435/471, 536/23.1, 536/23.2, 536/23.4, 536/23.7

US-CL-CURRENT: [514/44](#); [435/252.3](#), [435/254.11](#), [435/320.1](#), [435/325](#), [435/455](#), [435/471](#), [435/69.1](#), [536/23.1](#), [536/23.2](#), [536/23.4](#), [536/23.7](#)

FIELD-OF-SEARCH: 435/69.1, 435/320.1, 435/252.3, 435/254.11, 435/325, 435/455, 435/471, 536/23.1, 536/23.2, 536/23.4, 536/23.7, 514/44

PRIOR-ART-DISCLOSED:

OTHER PUBLICATIONS

El-Sherbeini, et al., "Staphylococcus aureus UDP-N-acetylmuramoyl-L-alanine: D-glutamate ligase (murD) gene, complete cds", GenBank Submission, Accession No. AF009671, Jul. 23, 1997.

Mengin-Lecreulx, et al., "Nucleotide sequence of the murD gene encoding the UDP-MurNAc-L-Ala-D-Glu synthetase of Escherichia coli", Nucleic Acids Research, vol. 18, No. 1, p. 183, Oxford University Press 1990.

El-Sherbeini, et al., "Cloning and expression of Staphylococcus aureus and Streptococcus pyogenes murD genes uridine diphosphate N-acetylmuramoyl-L-alanine:D-glutamate ligases", Gene, vol. 210, pp. 117-125, (1998).

Pucci, et al., "Staphylococcus aureus strain ATCC 8325-4 cell wall/cell division gene cluster, yllB, yllC, yllD, pbpA, murD, divlB, ftsA and ftsZ genes, complete cds.", GenBank Submission, Accession No. U94706, Jun. 3, 1997.

Pucci, et al., "Identification and Characterization of Cell Wall-Cell Division Gene Clusters in Pathogenic Gram-Positive Cocci", Journal of Bacteriology, vol. 179, No. 17, pp. 5632-5635, Sep. 1997.

Pucci, et al., Identification of bacterial UDP-N-acetyl muramyl-L-alanine: D-glutamate ligases. Abstracts of the 97th General Meeting of the American Society for Microbiology, May, 04-08, 1997, p. 360, abstract K-108.

ART-UNIT: 162

PRIMARY-EXAMINER: Priebe; Scott D.

ATTY-AGENT-FIRM: Gimmi; Edward R. Deibert; Thomas S. King; William T.

ABSTRACT:

The invention provides murD polypeptides and polynucleotides encoding murD polypeptides and methods for producing such polypeptides by recombinant techniques. Also provided are methods for utilizing murD polypeptides to screen for antibacterial compounds.

17 Claims, 0 Drawing figures

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L3: Entry 2 of 2

File: USPT

Apr 6, 1999

DOCUMENT-IDENTIFIER: US 5891621 A

TITLE: Metabolic pathway assay

Detailed Description Text (45):

In one embodiment of the present invention, the known gene sequences for murC, murD, murE and murF are used to synthesize their enzyme products: UDP-N-acetylmuramoyl-L-alanine ligase; UDP-N-acetylmuramoyl-L-alanine: D-glutamate ligase; UDP-N-acetylmuramoyl-L-alanyl-D-glutamate: meso-2,6-diaminopimelate ligase; and UDP-N-acetylmuramoyl-L-alanyl-D-glutamyl-meso-2,6-diaminopimoyl-D-alanine-D-alanine synthase, respectively. The genes are cloned by PCR and expressed using a modified commercially available GST or MBP fusion expression vector, such as pGEX-KT or pMal-C. The expression vector modification with respect to MBP fusions is detailed in Reference Example 3. The GST-fusion expression vectors are referenced under Hakes, D. J. and Dixon, J. E. "New Vectors for High Level Expression of Recombinant Proteins in Bacteria" 202 Analytical Biochemistry 293-298 (1992). The protein expression is detailed in Reference Example 5. The expressed enzymes are purified by affinity chromatography specific to the fusion protein used in the expression or any other suitable purification method. For example, GST fusions bind to glutathione agarose columns eluted with glutathione. MBP fusions bind to amylose columns eluted with maltose. The enzyme is cleaved from the purified fusion protein by incubating with thrombin. The contaminating GST or MBP can be removed by passage of the thrombin cleavage reaction mixture through the glutathione or amylose column one more time. The free enzyme passes through without binding to the column, while the GST or MBP will specifically stick to the column. The purification of the enzyme products of murC, murD, murE and murF is detailed in Reference Examples 6-9. In one embodiment of the present invention, the free mur enzymes are used for pathway assay. In another embodiment of the present invention, GST or MBP fusions of the mur enzymes are used for pathway assay. In an embodiment of the invention, MBP fusions are utilized.

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L4: Entry 1 of 15

File: USPT

Jan 4, 2005

US-PAT-NO: 6838255

DOCUMENT-IDENTIFIER: US 6838255 B1

TITLE: MurC

DATE-ISSUED: January 4, 2005

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Burnham; Martin Karl Russell	Norristown	PA		
Wallis; Nicola Gail	Wayne	PA		

ASSIGNEE-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY	TYPE CODE
SmithKline Beecham Corporation	Philadelphia	PA			02
SmithKline Beecham plc	Brentford			GB	03

APPL-NO: 09/ 103287 [PALM]

DATE FILED: June 23, 1998

PARENT-CASE:

RELATED APPLICATIONS This application claims benefit to U.S. Provisional Patent Application No. 60/052,720, filed Jul. 3, 1997.

INT-CL: [07] C12 P 21/02

US-CL-ISSUED: 435/69.1; 435/320.1, 435/252.3, 435/252.33, 536/23.7

US-CL-CURRENT: 435/69.1; 435/252.3, 435/252.33, 435/320.1, 536/23.7

FIELD-OF-SEARCH: 536/23.7, 536/23.4, 435/6, 435/455, 435/325, 435/252.3-257.35, 435/320.1, 435/69.1, 435/64.1

PRIOR-ART-DISCLOSED:

U.S. PATENT DOCUMENTS

Search Selected

Search ALL

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	PAT-NO	ISSUE-DATE	PATENTEE-NAME	US-CL
<input type="checkbox"/>	<u>6593114</u>	July 2003	Kunsch et al.	435/91.41
<input type="checkbox"/>	<u>6737248</u>	May 2004	Kunsch et al.	435/69.1

FOREIGN PATENT DOCUMENTS

FOREIGN-PAT-NO
0786519

PUBN-DATE
July 1997

COUNTRY
EP

US-CL

OTHER PUBLICATIONS

Varon, et al, Mol. Microbiol. 20: 339 (1996).
Critical Synergy: The Biotechnology Industry and Intellectual Property Protection,
Biotechnology Industry Organization, Washington, D.C., 1994, pp. 75 and 100-107.*
Genbank Submission, Accession No. AF034076, Direct Submission.
Swissprot Submission, Accession No. 030211, Direct Submission.

ART-UNIT: 1631

PRIMARY-EXAMINER: Martinell; James

ATTY-AGENT-FIRM: Fedon; Jason C. Gimmi; Edward R.

ABSTRACT:

The invention provides MurC polypeptides and polynucleotides encoding MurC polypeptides and methods for producing such polypeptides by recombinant techniques. Also provided are methods for utilizing MurC polypeptides to screen for antibacterial compounds.

14 Claims, 0 Drawing figures

First Hit Fwd Refs

L4: Entry 1 of 15

File: USPT

Jan 4, 2005

US-PAT-NO: 6838255

DOCUMENT-IDENTIFIER: US 6838255 B1

TITLE: MurC

DATE-ISSUED: January 4, 2005

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Burnham; Martin Karl Russell	Norristown	PA		
Wallis; Nicola Gail	Wayne	PA		

US-CL-CURRENT: 435/69.1; 435/252.3, 435/252.33, 435/320.1, 536/23.7

CLAIMS:

What is claimed is:

1. An isolated polynucleotide segment comprising a first polynucleotide sequence or the full complement of the entire length of the first polynucleotide sequence, wherein the first polynucleotide sequence encodes a polypeptide comprising the amino acid sequence of SEQ ID NO: 2.
2. A vector comprising the isolated polynucleotide segment of claim 1.
3. An isolated host cell comprising the vector of claim 2.
4. A process for producing a polypeptide comprising the step of culturing the host cell of claim 3 under conditions sufficient for the production of the polypeptide, wherein the polypeptide is encoded by the first polynucleotide sequence.
5. The isolated polynucleotide of claim 1 encoding a fusion polypeptide, wherein the first polynucleotide sequence encodes part of the fusion polypeptide.
6. An isolated polynucleotide segment comprising a first polynucleotide sequence or the full complement of the entire length of the first polynucleotide sequence, wherein the first polynucleotide sequence encodes a polypeptide consisting of the amino acid sequence of SEQ ID NO:2.
7. A vector comprising the isolated polynucleotide segment of claim 6.
8. An isolated host cell comprising the vector of claim 7.
9. A process for producing a polypeptide comprising the step of culturing the host cell of claim 8 under conditions sufficient for the production of the polypeptide, wherein the polypeptide is encoded by the first polynucleotide

sequence.

10. An isolated polynucleotide segment comprising a first polynucleotide sequence or the full complement of the entire length of the first polynucleotide sequence wherein the first polynucleotide sequence comprises SEQ ID NO:1.

11. A vector comprising the isolated polynucleotide segment of claim 10.

12. An isolated host cell comprising the vector of claim 11.

13. A process for producing a polypeptide comprising the step of culturing the host cell of claim 12, under conditions sufficient for the production of the polypeptide, wherein the polypeptide is encoded by the first polynucleotide sequence.

14. The isolated polynucleotide of claim 10 encoding a fusion polypeptide, wherein the first polynucleotide sequence encodes part of the fusion polypeptide.